

TABLE S1: The oligonucleotide primers used for the real-time qPCR

| Genes | Primer sequences |
|---------------|-----------------------------|
| <i>efl1α</i> | 5'-CGTGGTAATGTGGCTGGAGA |
| | 5'-CTGAGCGTTGAAGTTGGCAG |
| <i>fbp1a</i> | 5'-ATCGTTAGAGCCAGATAGAC |
| | 5'-TGTAGATGGCAAAGATGGTCC |
| <i>gclm</i> | 5'-ATCCATCAGAAGTGCCTAG |
| | 5'-AGTGTCTGGCTTCACCCTC |
| <i>gsr</i> | 5'-CAGATTACAGGAGCAACCTG |
| | 5'-TGACTTCAACTGTGGGTTTAG |
| <i>gsto2</i> | 5'-ATGGCTTCATCTCCAAAATGC |
| | 5'-AGGGCAGAACATCTCATGCTGTAG |
| <i>gstp1</i> | 5'-CAACGCCATGCTGAGACATC |
| | 5'-GAAGATCTTCAACGCCGTG |
| <i>keap1a</i> | 5'-ATACCAACCAGACACCAACAC |
| | 5'-GGTTGTCCATCATAGCCTCC |
| <i>nrf2a</i> | 5'-ATGTCTAAATGCAGCCAAGCC |
| | 5'-CGGTAGCTGAAGTCGAACAC |
| <i>pck1</i> | 5'-GGAGAACAGCACCATCCTCAG |
| | 5'-AGTCGGTGTGGAACCGTG |
| <i>pcxb</i> | 5'-AGATGGGGATAAAGTGGAG |
| | 5'-TCTGAAGGCATGAGATAGGAG |
| <i>pgd</i> | 5'-TCCAATACGGCACACCTGTC |
| | 5'-AGGCTCTTACTGGCCTGAAC |
| <i>psma3</i> | 5'-ACGGAAGGGTATTTCAGGTTG |
| | 5'-TCATACAGCTTGGACAGTACC |
| <i>psma5</i> | 5'-AGTAAACACTTCTCACCAGAAG |
| | 5'-TCTCCACAGCAAGACAAACTC |
| <i>psmb7</i> | 5'-GAGCTTCATTCTCTGTCCACC |
| | 5'-GCAATCGACACCTCCAAGAAC |
| <i>sqstm1</i> | 5'-TTGGCTTTGTGAAGGATGAC |
| | 5'-TGTAGTGAACGGAAACCCAGG |
| <i>taldo1</i> | 5'-TCTGTATCACCGGACAAACG |
| | 5'-TCCTGAGGCTGTACTCTTC |

TABLE S2: The oligonucleotide primers used for plasmid construction

| Plasmids | Primer sequences |
|----------|------------------------------------|
| pKSpsma3 | 5'-GGGGGATCCTGATCGCAGTGACTTAAAATG |
| | 5'-GGGCTCGAGACATGTTGCCTCATCAGAGTC |
| pKSpsma5 | 5'-GGGAAATTCTCCACTGCAGAAGATTGCG |
| | 5'-GGGCTCGAGAGCACGTTAGATATCCTTGATG |
| pKSpsmb7 | 5'-GGGGGATCCAGACCGCAGAACATGGCGAC |
| | 5'-GGGCTCGAGTCTGTTGGCAATGTCGTGAG |
| pKSpgd | 5'-GGGGGATCCAGCAGTCAAACCCCTCATCG |
| | 5'-GGGCTCGAGCATTCCCTGTGCCCTCTG |
| pKSfbp1a | 5'-GGGGGATCCGATGGTCATCCAACATTGAC |
| | 5'-GGGCTCGAGGGCATCTTATTCTGGAGTGTC |

TABLE S3: The genes induced by the overexpression of *nrf2a* in zebrafish.

| Ratio | Gene symbol in ZFIN (or [19]) | Gene product | Refs. | |
|-------|----------------------------------|--|------------|---------------------|
| | | | microarray | confirmed by PCR |
| 9.96 | <i>gclm</i> | glutamate-cysteine ligase, modifier subunit | [19,20] | [19] |
| 7.34 | <i>hmxox1a</i> | heme oxygenase 1a | – | [19] |
| 6.53 | <i>pimr138</i> | Pim proto-oncogene, serine/threonine kinase, related 138 | – | – |
| 5.81 | <i>aifm4</i> | apoptosis-inducing factor, mitochondrion-associated, 4 | [19,20] | – |
| 5.71 | <i>gstp1</i> | glutathione S-transferase pi 1 | [19] | [11,16,17,19] |
| 4.50 | <i>txn</i> | thioredoxin | [19,20] | [16] |
| 4.02 | <i>zp2l1</i> | zona pellucida glycoprotein 2, like 1 | – | – |
| 3.83 | <i>zgc:92066 (fthl)</i> | ferritin heavy chain like | [19,20] | [19] |
| 3.39 | <i>tex2</i> | testis expressed 2 | [19] | – |
| 3.21 | <i>gsto2</i> | glutathione S-transferase omega 2 | [19] | – |
| 2.98 | <i>gclc</i> | glutamate-cysteine ligase, catalytic subunit | [20] | [16,17,19] |
| 2.84 | <i>pcxb</i> | pyruvate carboxylase b | [19] | – |
| 2.70 | <i>exorh</i> | extra-ocular rhodopsin | – | – |
| 2.65 | <i>zfand2a</i> | zinc finger, AN1-type domain 2A | – | – |
| 2.49 | <i>ugdh</i> | UDP-glucose 6-dehydrogenase | [19] | – |
| 2.47 | <i>mgst3b</i> | microsomal glutathione S-transferase 3b | [19,20] | [19] |
| 2.37 | <i>sepw2b</i> | selenoprotein W, 2b | [19] | [19] |
| 2.36 | <i>pisd</i> | phosphatidylserine decarboxylase | – | – |
| 2.33 | <i>lgals2a</i> | lectin, galactoside-binding, soluble, 2a | – | – |
| 2.32 | <i>cbr1l</i> | carbonyl reductase 1-like | [19] | – |
| 2.25 | <i>lrrc6</i> | leucine rich repeat containing 6 | – | – |
| 2.25 | <i>tjp2b</i> | tight junction protein 2b (zona occludens 2) | – | – |
| 2.13 | <i>sqndl</i> | sulfide quinone reductase-like (yeast) | [19] | – |
| 2.13 | <i>spata6l</i> | spermatogenesis associated 6-like | – | – |
| 2.11 | <i>sec23b</i> | Sec23 homolog B, COPII coat complex component | – | – |
| 2.11 | <i>slc40a1</i> | solute carrier family 40 (iron-regulated transporter), member 1 | – | – |
| 2.09 | <i>rab43</i> | RAB43, member RAS oncogene family | [19] | – |
| 2.08 | <i>sqstm1</i> | sequestosome 1 | [20] | – |
| 2.07 | <i>lmo7a</i> | LIM domain 7a | – | – |
| 2.06 | <i>hsp90aa1.1</i> | heat shock protein 90, alpha (cytosolic), class A member 1, tandem duplicate 1 | – | – |
| 2.00 | <i>keap1a</i> | kelch-like ECH-associated protein 1a | [20] | – |
| 2.00 | <i>fam13b</i> | family with sequence similarity 13, member B | – | – |
| 1.94 | <i>tradd</i> | tnfrsf1a-associated via death domain | – | – |
| 1.94 | <i>ctsk</i> | cathepsin K | – | – |
| 1.94 | <i>selp</i> | selectin P | – | – |
| 1.92 | <i>tpk1</i> | thiamin pyrophosphokinase 1 | – | – |
| 1.91 | <i>rsrp1</i> | arginine-serine-rich protein 1 | [20] | – |
| 1.87 | <i>gstal (gstal)</i> | glutathione S-transferase, alpha tandem duplicate 1 | [19] | [19] |
| 1.87 | <i>epoa</i> | erythropoietin a | – | – |
| 1.85 | <i>pfas</i> | phosphoribosylformylglycinamide synthase | – | – |
| 1.85 | <i>setdb1a</i> | SET domain, bifurcated 1a | – | – |

Gray highlighting indicate genes that were also identified in the previous microarray analyses using DEM- or tBHQ-treated larvae [19,20].

TABLE S3: The genes induced by the overexpression of *nrf2a* in zebrafish. (continued)

| Ratio | Gene symbol in ZFIN (or [19]) | Gene product | Refs. | |
|-------|----------------------------------|--|------------|---------------------|
| | | | microarray | confirmed by PCR |
| 1.84 | <i>apoal1a</i> | apolipoprotein A-Ia | – | – |
| 1.83 | <i>gpx1b</i> | glutathione peroxidase 1b | – | – |
| 1.80 | <i>pik3c2a</i> | phosphatidylinositol-4-phosphate 3-kinase, catalytic subunit type 2 alpha | – | – |
| 1.80 | <i>psma3</i> | proteasome subunit alpha 3 | – | – |
| 1.78 | <i>cyp2r1</i> | cytochrome P450, family 2, subfamily R, polypeptide 1 | – | – |
| 1.78 | <i>si:dkey-33c12.4</i> | tetratricopeptide repeat domain 31 (ttc31) (ENSDARG00000057890) | – | – |
| 1.78 | <i>esrp1</i> | epithelial splicing regulatory protein 1 | – | – |
| 1.78 | <i>si:ch211-95j8.2</i> | claudin 23 like (ENSDARG00000028096) | – | – |
| 1.78 | <i>coa3a</i> | cytochrome C oxidase assembly factor 3a | – | – |
| 1.72 | <i>epn2</i> | epsin 2 | – | – |
| 1.72 | <i>brf1a</i> | BRF1, RNA polymerase III transcription initiation factor a | – | – |
| 1.71 | <i>col4a1</i> | collagen, type IV, alpha 1 | – | – |
| 1.71 | <i>ndrg3a</i> | ndrg family member 3a | – | – |
| 1.69 | <i>tp53bp2a</i> | tumor protein p53 binding protein, 2a | – | – |
| 1.68 | <i>cxcr4b</i> | chemokine (C-X-C motif), receptor 4b | – | – |
| 1.68 | <i>phyh</i> | phytanoyl-CoA 2-hydroxylase | – | – |
| 1.68 | <i>nudc</i> | nudC nuclear distribution protein | – | – |
| 1.68 | <i>gldc</i> | glycine dehydrogenase (decarboxylating) | – | – |
| 1.68 | <i>taldo1</i> | transaldolase 1 | – | – |
| 1.67 | <i>si:dkey-122c11.8</i> | novel gene, “zinc finger protein” | – | – |
| 1.67 | <i>anxa1a</i> | annexin A1a | – | – |
| 1.66 | <i>zgc:163022 (frs1c)</i> | ferric-chelate reductase like (CABZ01041812.1) | [19] | [19] |
| 1.66 | <i>dnase1l3</i> | deoxyribonuclease I-like 3 | – | – |
| 1.65 | <i>atp2b2</i> | ATPase, Ca ⁺⁺ transporting, plasma membrane 2 | – | – |
| 1.63 | <i>rbm24a</i> | RNA binding motif protein 24a | – | – |
| 1.62 | <i>uqcrq</i> | ubiquinol-cytochrome c reductase, complexIII subunit VII | – | – |
| 1.62 | <i>rab3il1</i> | RAB3A interacting protein (rabin3)-like 1 | – | – |
| 1.62 | <i>sst6</i> | somatostatin 6 | – | – |
| 1.62 | <i>myh9a</i> | myosin, heavy chain 9a, non-muscle | – | – |
| 1.61 | <i>kpnal1</i> | karyopherin alpha 1 (importin alpha 5) | – | – |
| 1.60 | <i>pck1</i> | phosphoenolpyruvate carboxykinase 1 (soluble) | [19] | – |
| 1.60 | <i>psma5</i> | proteasome subunit alpha 5 | – | – |
| 1.60 | <i>prdx5</i> | peroxiredoxin 5 | – | – |
| 1.60 | <i>gpx1a</i> | glutathione peroxidase 1a | – | – |
| 1.59 | <i>lmna</i> | lamin A | – | – |
| 1.57 | <i>tmem106a</i> | transmembrane protein 106A | – | – |
| 1.57 | <i>gramd1c</i> | GRAM domain containing 1c | – | – |
| 1.57 | <i>si:ch211-106h4.6</i> | novel gene, “plexin domain containing like” | – | – |
| 1.57 | <i>hoxb6b</i> | homeobox B6b | – | – |
| 1.56 | <i>srprb</i> | signal recognition particle receptor, B subunit | – | – |

Gray highlighting indicate genes that were also identified in the previous microarray analyses using DEM- or tBHQ-treated larvae [19,20].

TABLE S3: The genes induced by the overexpression of *nrf2a* in zebrafish. (continued)

| Ratio | Gene symbol in ZFIN (or [19]) | Gene product | Refs. | |
|-------|----------------------------------|---|------------|---------------------|
| | | | microarray | confirmed by PCR |
| 1.55 | <i>slx4</i> | SLX4 structure-specific endonuclease subunit homolog (<i>S. cerevisiae</i>) | — | — |
| 1.55 | <i>gsr</i> | glutathione reductase | [20] | — |
| 1.55 | <i>ftr82</i> | fin TRIM family, member 82 | — | — |
| 1.55 | <i>map1b</i> | microtubule-associated protein 1B | — | — |
| 1.55 | <i>prkcdb</i> | protein kinase C, delta b | [19] | — |
| 1.55 | <i>pth1ra</i> | parathyroid hormone 1 receptor a | — | — |
| 1.55 | <i>calm3a</i> | calmodulin 3a (phosphorylase kinase, delta) | — | — |
| 1.55 | <i>gatm</i> | glycine amidinotransferase (L-arginine:glycine amidinotransferase) | — | — |
| 1.54 | <i>smc5</i> | structural maintenance of chromosomes 5 | — | — |
| 1.54 | <i>caska</i> | calcium/calmodulin-dependent serine protein kinase a | — | — |
| 1.54 | <i>opn1mw1</i> | opsin 1 (cone pigments), medium-wave-sensitive, 1 | — | — |
| 1.54 | <i>lcp1</i> | lymphocyte cytosolic protein 1 (L-plastin) | — | — |
| 1.54 | <i>aoc2</i> | amine oxidase, copper containing 2 | [19] | — |
| 1.54 | <i>epb41l5</i> | erythrocyte membrane protein band 4.1 like 5 | — | — |
| 1.54 | <i>tacc2</i> | transforming, acidic coiled-coil containing protein 2 | — | — |
| 1.53 | <i>rbm24a</i> | RNA binding motif protein 24a | — | — |
| 1.52 | <i>prkacab</i> | protein kinase, cAMP-dependent, catalytic, alpha, genome duplicate b | — | — |
| 1.52 | <i>runx1t1</i> | runt-related transcription factor 1; translocated to, 1(cyclin D-related) | — | — |
| 1.52 | <i>gmeb1</i> | glucocorticoid modulatory element binding protein 1 | — | — |
| 1.52 | <i>ctsh</i> | cathepsin H | — | — |
| 1.52 | <i>sult6b1</i> | sulfotransferase family, cytosolic, 6b, member 1 | [19] | — |
| 1.52 | <i>fbp1a</i> | fructose-1,6-bisphosphatase 1a | — | — |
| 1.51 | <i>pgd</i> | phosphogluconate dehydrogenase | [19,20] | — |
| 1.51 | <i>slc4a2a</i> | solute carrier family 4 (anion exchanger), member 2a | — | — |
| 1.50 | <i>ppiaa</i> | peptidylprolyl isomerase Aa (cyclophilin A) | — | — |
| 1.50 | <i>zyx</i> | zyxin | — | — |
| 1.50 | <i>rabif</i> | RAB interacting factor | — | — |
| 1.50 | <i>acss1</i> | acyl-CoA synthetase short-chain family member 1 | — | — |
| 1.50 | <i>si:ch211-87h14.3</i> | chromosome 19 open reading frame 60 (C19orf60) | — | — |
| 1.50 | <i>psmb7</i> | proteasome subunit beta 7 | — | — |
| 1.50 | <i>penkb</i> | proenkephalin b | — | — |
| 1.50 | <i>sepsecs</i> | Sep (O-phosphoserine) tRNA: Sec (selenocysteine) tRNA synthase | — | — |
| 1.50 | <i>atp6v0a1a</i> | ATPase, H ⁺ transporting, lysosomal V0 subunit a1a | — | — |
| 1.50 | <i>prdx1</i> | peroxiredoxin 1 | [19,20] | [16,17,19] |

Gray highlighting indicate genes that were also identified in the previous microarray analyses using DEM- or tBHQ-treated larvae [19,20].

TABLE S4: The genes repressed by the overexpression of *nrf2a* in zebrafish.

| Ratio | Gene symbol in ZFIN | Gene product |
|-------|---------------------|---|
| 0.31 | <i>ccnc</i> | cyclin C |
| 0.40 | <i>prkacbb</i> | protein kinase, cAMP-dependent, catalytic, beta b |
| 0.48 | <i>dynll2a</i> | dynein, light chain, LC8-type 2a |
| 0.51 | <i>slco1f1</i> | solute carrier organic anion transporter family, member 1F1 |
| 0.54 | <i>ipo7</i> | importin 7 |
| 0.55 | <i>tmem135</i> | transmembrane protein 135 |
| 0.56 | <i>hesx1</i> | HESX homeobox 1 |
| 0.57 | <i>vps51</i> | vacuolar protein sorting 51 homolog (<i>S. cerevisiae</i>) |
| 0.57 | <i>cotl1</i> | coactosin-like F-actin binding protein 1 |
| 0.58 | <i>sparc</i> | secreted protein, acidic, cysteine-rich (osteonectin) |
| 0.58 | <i>phgdh</i> | phosphoglycerate dehydrogenase |
| 0.59 | <i>sltm</i> | SAFB-like, transcription modulator |
| 0.60 | <i>pfkpb</i> | phosphofructokinase, platelet b |
| 0.60 | <i>taf15</i> | TAF15 RNA polymerase II, TATA box binding protein (TBP)-associated factor |
| 0.60 | <i>kpnb3</i> | karyopherin (importin) beta 3 |
| 0.61 | <i>ncl</i> | nucleolin |
| 0.61 | <i>efemp2a</i> | EGF containing fibulin-like extracellular matrix protein 2a |
| 0.62 | <i>srsf5a</i> | serine/arginine-rich splicing factor 5a |
| 0.62 | <i>rnf19a</i> | ring finger protein 19A, RBR E3 ubiquitin protein ligase |
| 0.62 | <i>cox15</i> | cytochrome c oxidase assembly homolog 15 (yeast) |
| 0.63 | <i>fbxo11a</i> | F-box protein 11a |
| 0.63 | <i>etflb</i> | eukaryotic translation termination factor 1b |
| 0.63 | <i>dmrt3a</i> | doublesex and mab-3 related transcription factor 3a |
| 0.63 | <i>anp32e</i> | acidic (leucine-rich) nuclear phosphoprotein 32 family, member E |
| 0.63 | <i>gnl3</i> | guanine nucleotide binding protein-like 3 (nucleolar) |
| 0.63 | <i>dennd4c</i> | DENN/MADD domain containing 4C |
| 0.64 | <i>src</i> | v-src avian sarcoma (Schmidt-Ruppin A-2) viral oncogene homolog |
| 0.64 | <i>anos1b</i> | anosmin 1b |
| 0.64 | <i>pfkfb4b</i> | 6-phosphofructo-2kinase/fructose-2,6-biphosphatase 4b |
| 0.64 | <i>gskip</i> | gsk3b interacting protein |
| 0.64 | <i>cirbp</i> | cold inducible RNA binding protein a |
| 0.64 | <i>sec14l1</i> | SEC14-like lipid binding 1 |
| 0.64 | <i>tcf15</i> | transcription factor 15 (basic helix-loop-helix) |
| 0.64 | <i>brd2b</i> | bromodomain containing 2b |
| 0.64 | <i>nudt3a</i> | nudix (nucleoside diphosphate linked moiety X)-type motif 3a |
| 0.65 | <i>atrx</i> | alpha thalassemia/mental retardation syndrome X-linked homolog (human) |
| 0.65 | <i>zgc:153953</i> | chromosome 1 open reading frame 106 (C1orf106) |
| 0.65 | <i>zgc:113452</i> | novel gene “zink finger protein” |
| 0.65 | <i>kctd4</i> | potassium channel tetramerization domain containing 4 |
| 0.65 | <i>fam60al</i> | family with sequence similarity 60, member A, like |
| 0.66 | <i>tle2</i> | transducing-like enhancer of split 2 (E(sp1) homolog, <i>Drosophila</i>) |
| 0.66 | <i>mcm6</i> | minichromosome maintenance complex component 6 |

TABLE S4: The genes repressed by the overexpression of *nrf2a* in zebrafish. (continued)

| Ratio | Gene symbol in ZFIN | Gene product |
|-------|---------------------|--|
| 0.66 | <i>gsna</i> | gelsolin a |
| 0.66 | <i>snapin</i> | SNAP-associated protein |
| 0.66 | <i>fscn1a</i> | fascin actin-bundling protein 1a |
| 0.66 | <i>hoxa10b</i> | homeobox A 10b |
| 0.67 | <i>bmpr1aa</i> | bone morphogenetic protein receptor, type IAa |
| 0.67 | <i>stk35</i> | serine/threonine kinase 35 |
| 0.67 | <i>lsm12a</i> | LSM 12 homolog a |
| 0.67 | <i>tfap2a</i> | transcription factor AP-2 alpha |
| 0.67 | <i>ptbp1a</i> | polypyrimidine tract binding protein 1a |
| 0.67 | <i>fzd8a</i> | frizzled class receptor 8a |
| 0.67 | <i>sema4c</i> | sema domain, immunoglobulin domain (Ig), transmembrane domain (TM) and short cytoplasmic domain (semaphorin), 4C |
| 0.67 | <i>lrwd1</i> | leucine-rich repeats and WD repeat domain containing 1 |
| 0.67 | <i>supt6h</i> | SPT6 homolog, histone chaperone |